

cells being modified by homologous recombination with the DNA fragment from (a),

- AA  
CONT.
- (d) culturing the cells from step (c),
  - (e) [selection of] selecting the cultured cells from step (d) for the absence of the homologous sequence and/or the presence of the heterologous sequence, [in particular the selectable marker,]
  - (f) [production of] producing chimeric non-human mammals from the cells from step (e) by injection of these cells in mammalian blastocysts [(preferably mouse blastocysts)], [transfer of] transferring the blastocysts into false-pregnant female mammals [(preferably mouse)] and [analysis of] analyzing the resulting offspring for a change of the NINTROX gene.

---

**REMARKS**

Entry of the foregoing amendments are respectfully requested.

Should the Examiner have any questions concerning the subject application, a telephone call to the undersigned would be appreciated.

Respectfully submitted,

BURNS, DOANE, SWECKER & MATHIS, L.L.P.

By: Teresa Stanek Rea  
Teresa Stanek Rea  
Registration No. 30,427

*for F.J. Den*  
Reg. No. 28,570

P.O. Box 1404  
Alexandria, Virginia 22313-1404  
(703) 836-6620

Date: December 22, 2000